

Supplementary information, Fig. S7. In this figure, different normal or cancer cell lines were treated with DM- α KG (15 mM) for 24 hours to indicate cell morphology, unless specially indicated otherwise.

(a) Comparison of different protein expression levels in pyroptosis-sensitive and pyroptosis-insensitive cancer cell lines. Protein expression levels were measured by western blot.

(b) Intracellular pH was determined in the different pyroptosis-sensitive and pyroptosis-insensitive cell lines.

(c, d) Different cancer (c) or non-cancer (d) cell lines were cultured in medium under different pH conditions as indicated for 24 hours. The pyroptotic features are shown.

(e) Effects of different pH conditions on DM- α KG induced pyroptosis in HeLa and B16 cells.

(f) Lactic acid decreased the intracellular pH in different cancer cell lines. Cells were treated with lactic acid (20 mM) for 0.5 hours.

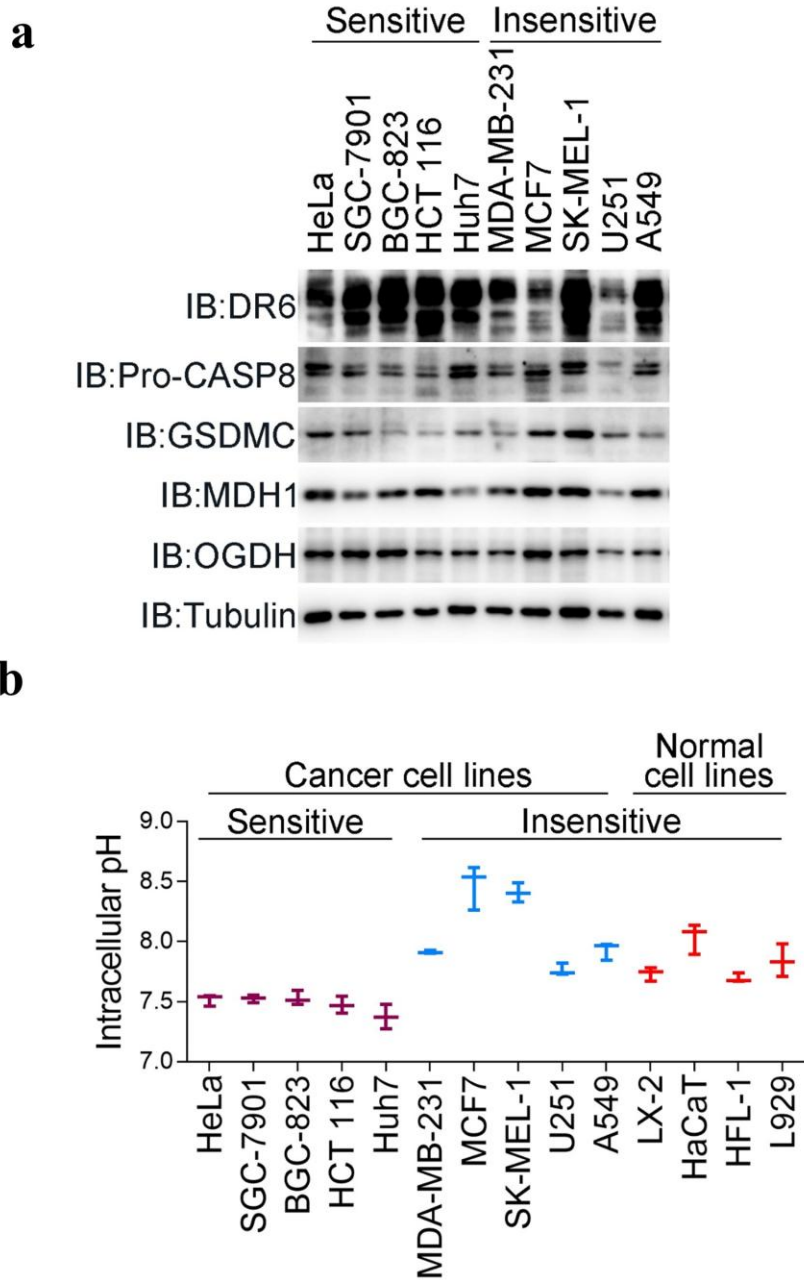
(g) Lactic acid induced the conversion of pyroptosis-insensitive cells to pyroptosis-sensitive cells. Cells were treated with lactic acid (20 mM) for 24 hours.

(h, i, j, k) Effect of phloretin (50 μ M) or Ionidamine (50 μ M) on DM- α KG induced pyroptosis (h, i), intracellular pH (j) and L-2HG levels (k). HeLa cells were pretreated

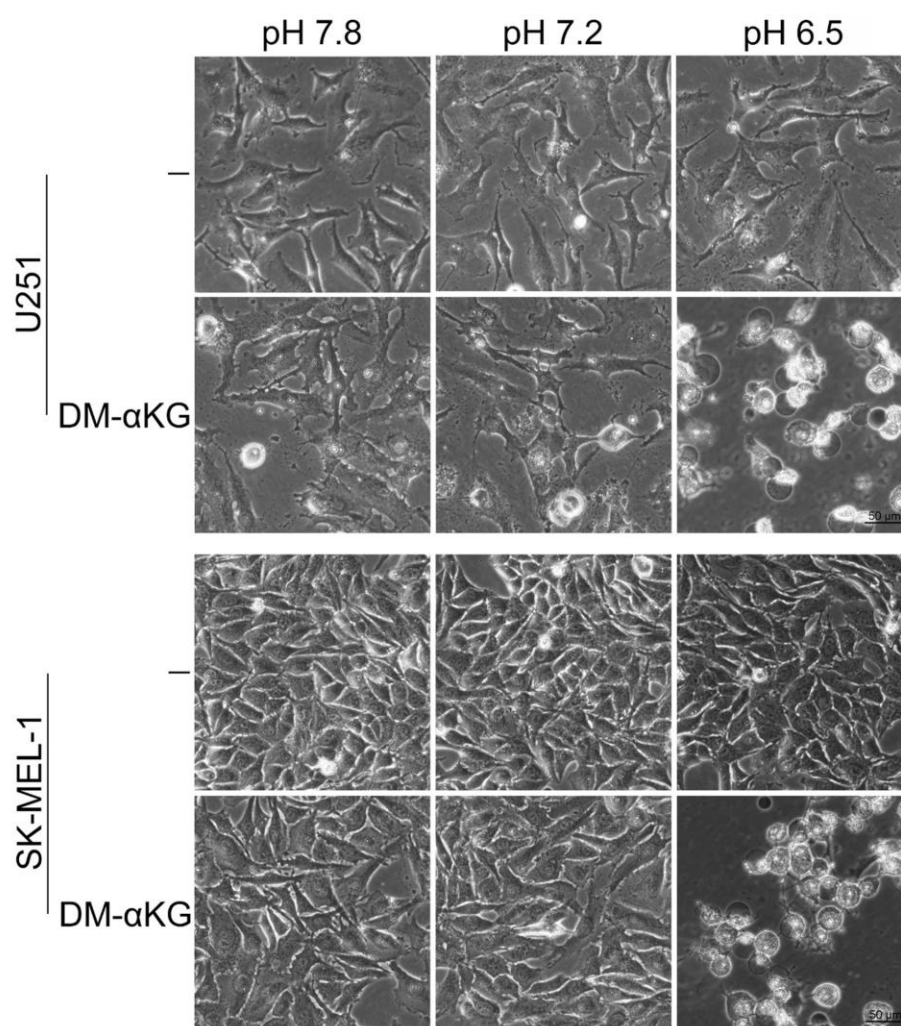
with phloretin or lonidamine for 2 hours.

Tubulin was used to determine the amount of loading proteins. All data are presented as the mean \pm SEM of two or three independent experiments. * $p<0.05$, ** $p<0.01$, *** $p<0.001$. The data were analyzed using two-tailed Student's t-test in (f, j) or two-way ANOVA followed by the Bonferroni test in (e, h, i, k).

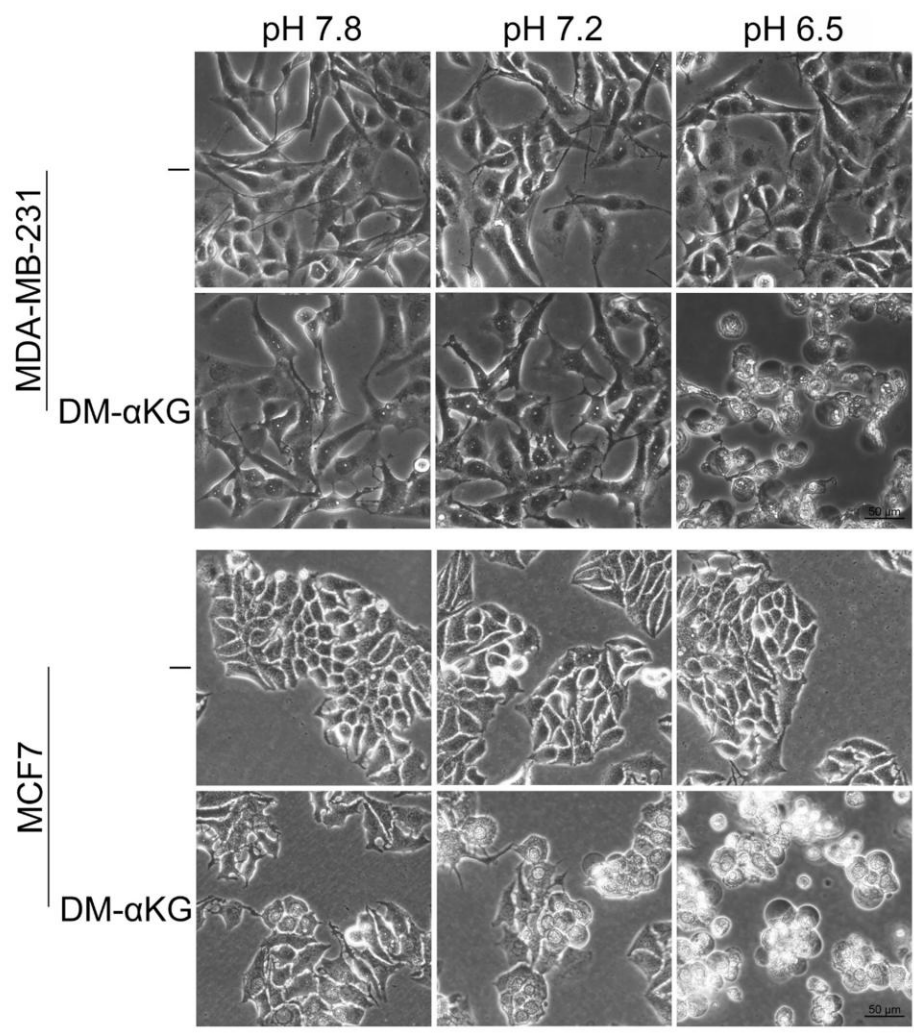
Supplementary information, Figure S7

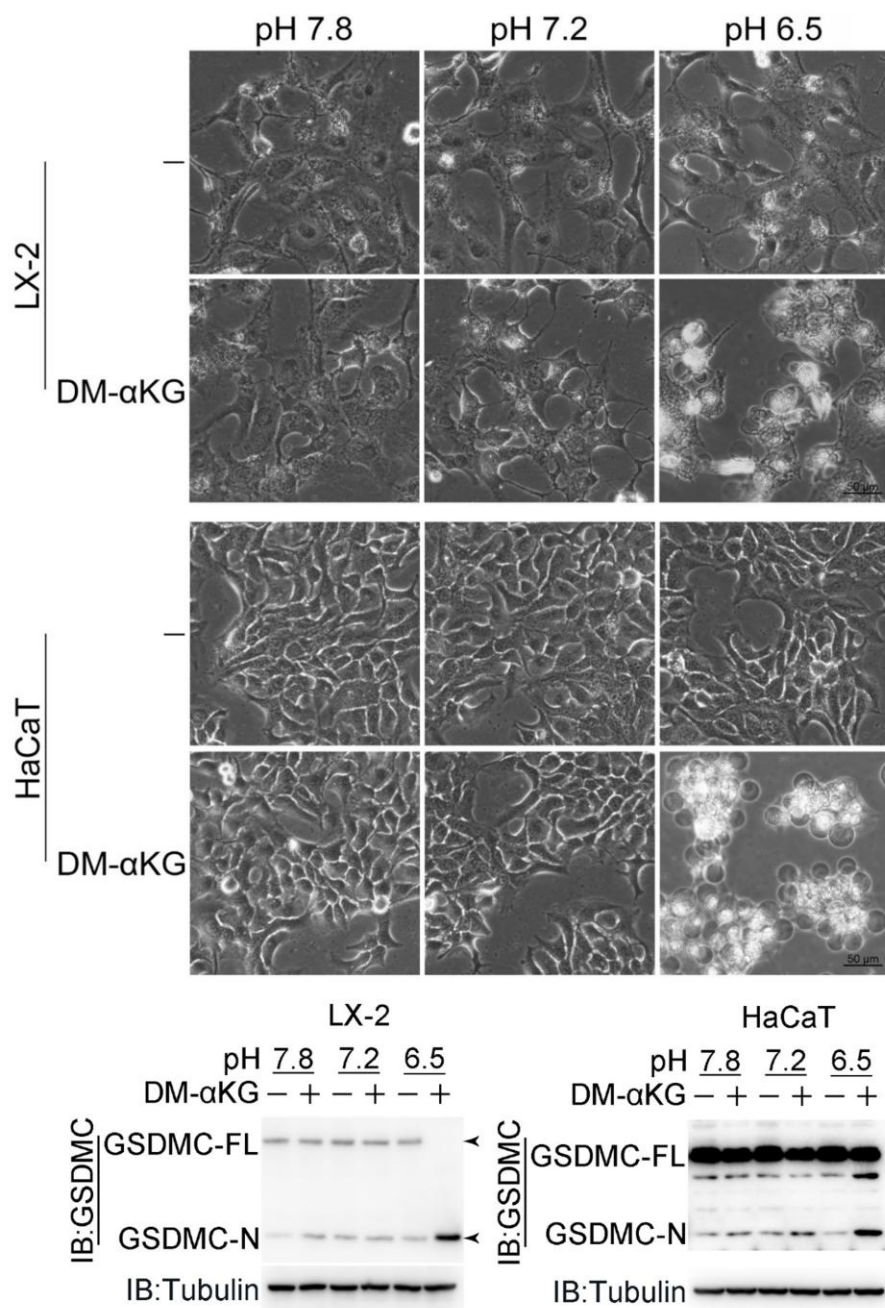


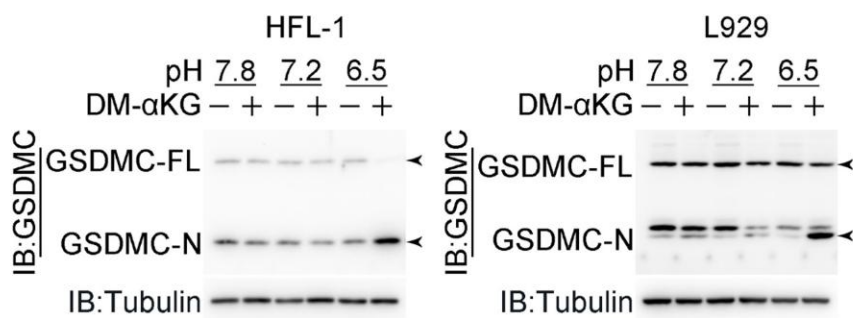
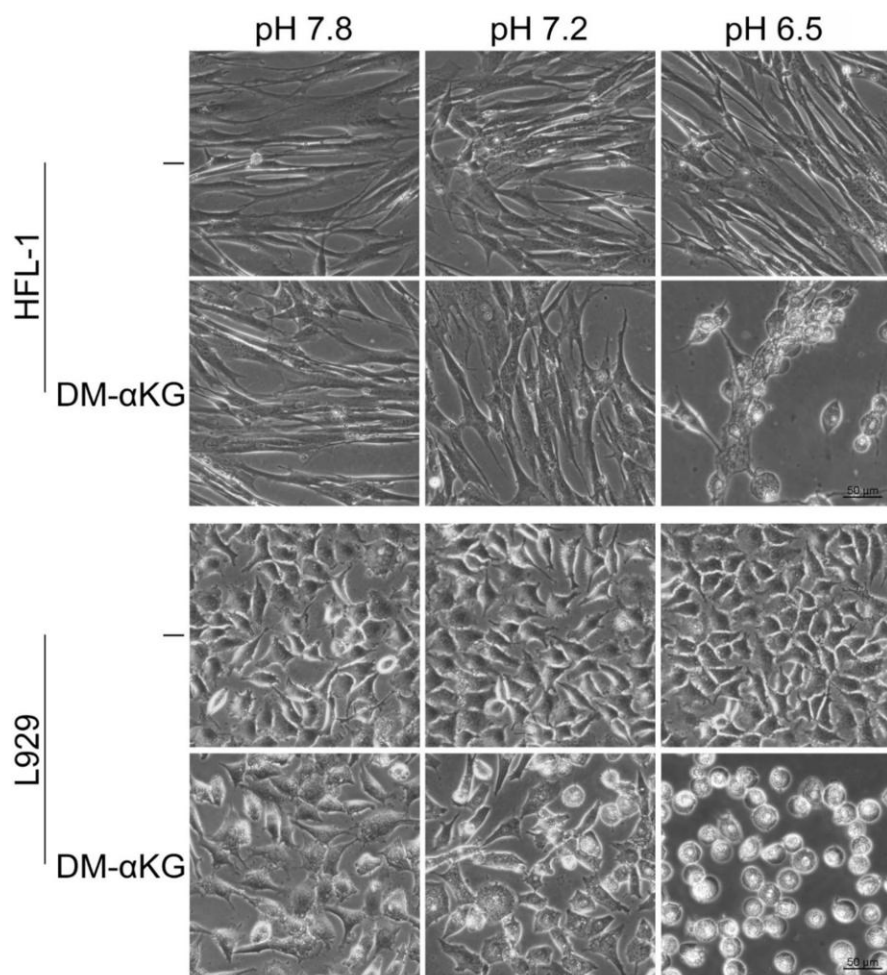
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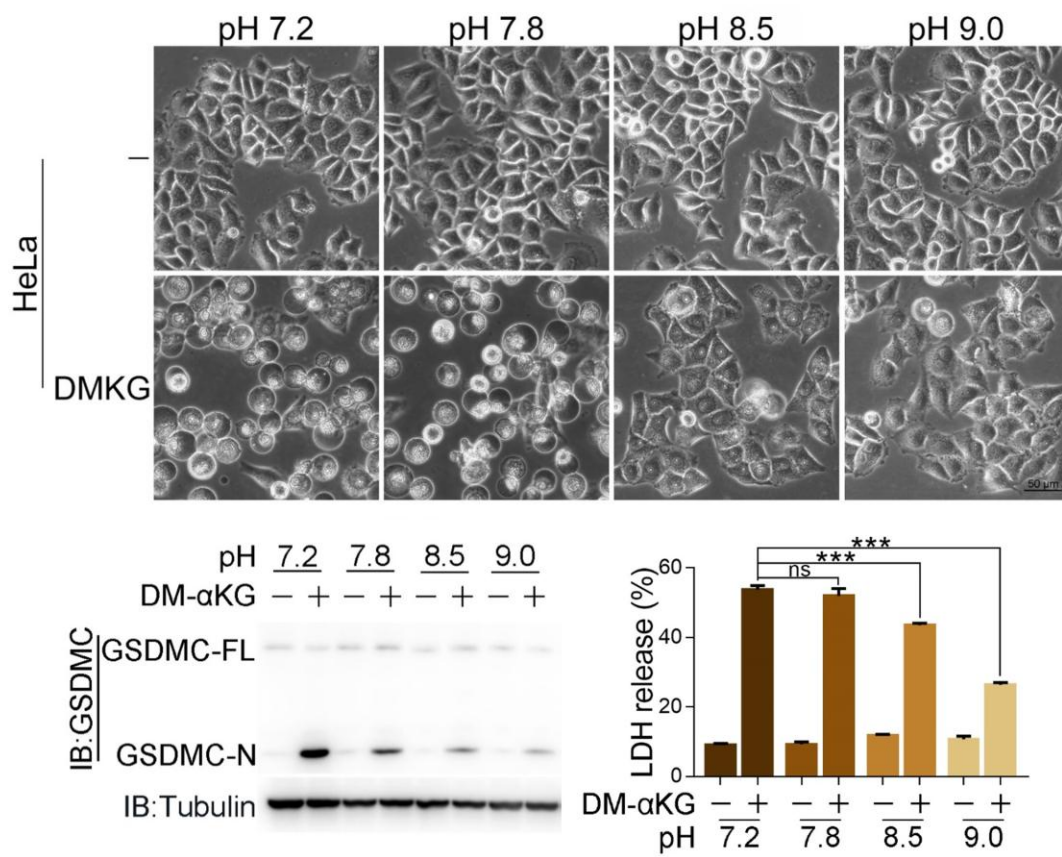
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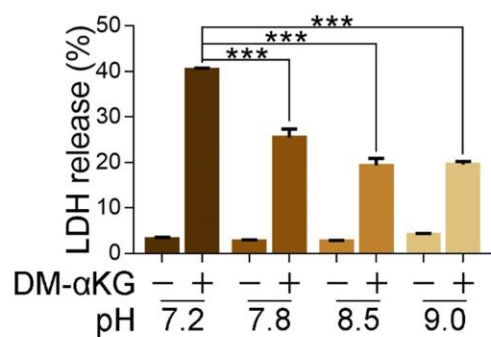
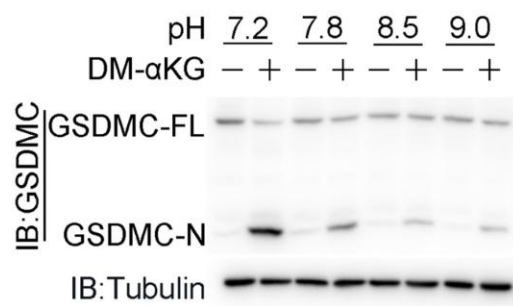
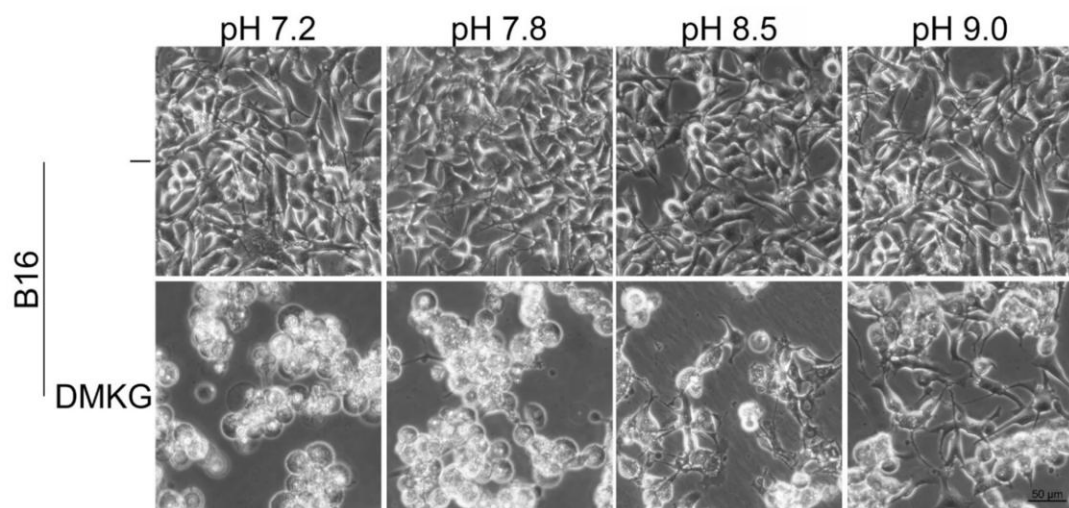
d**(Continue)**

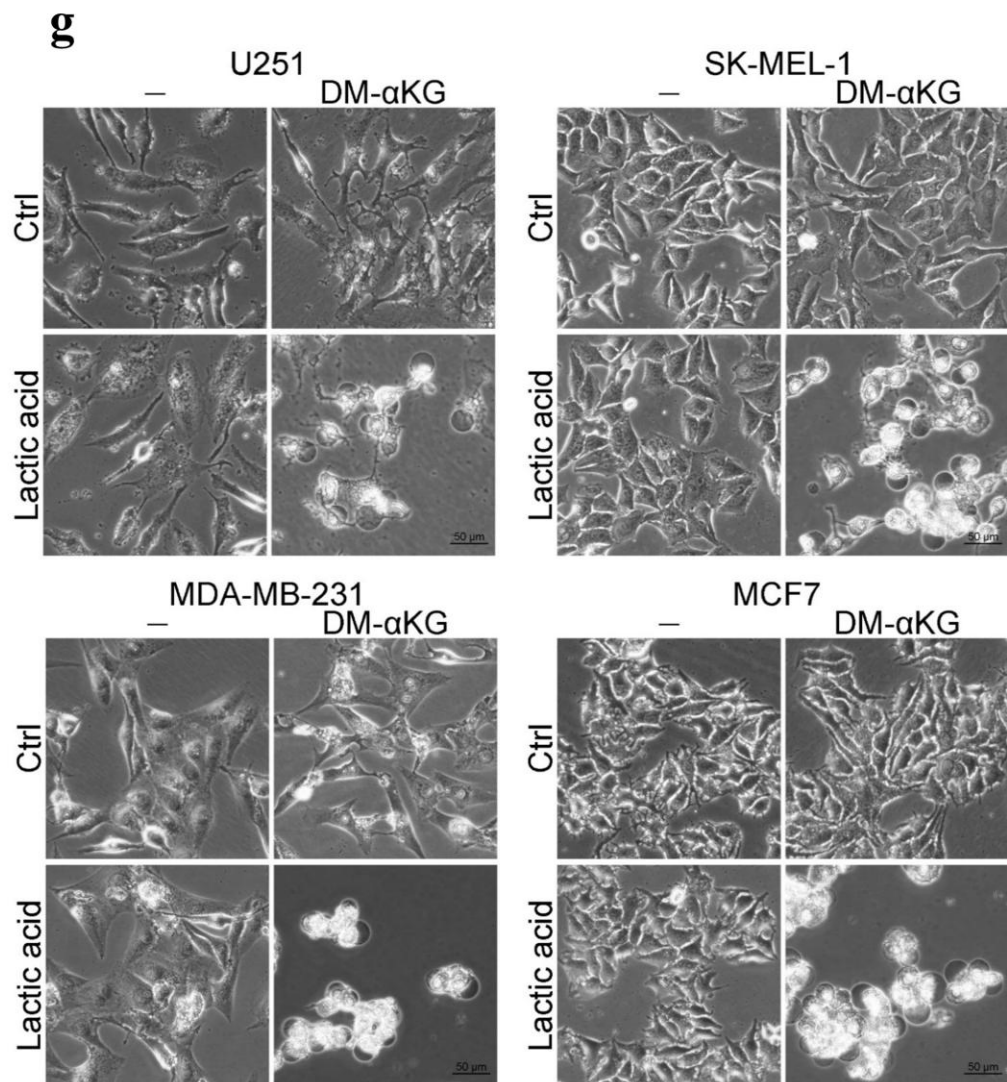
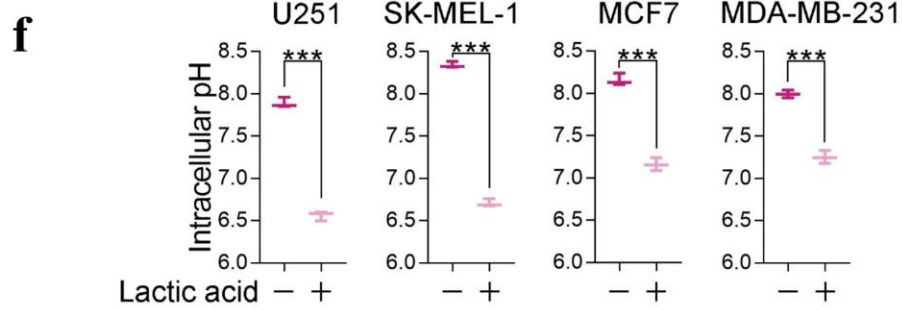


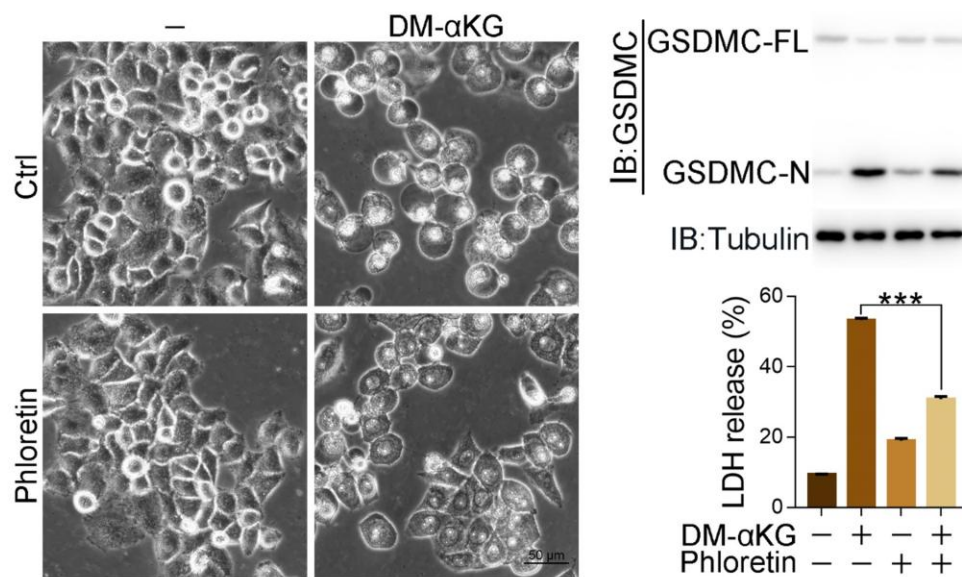
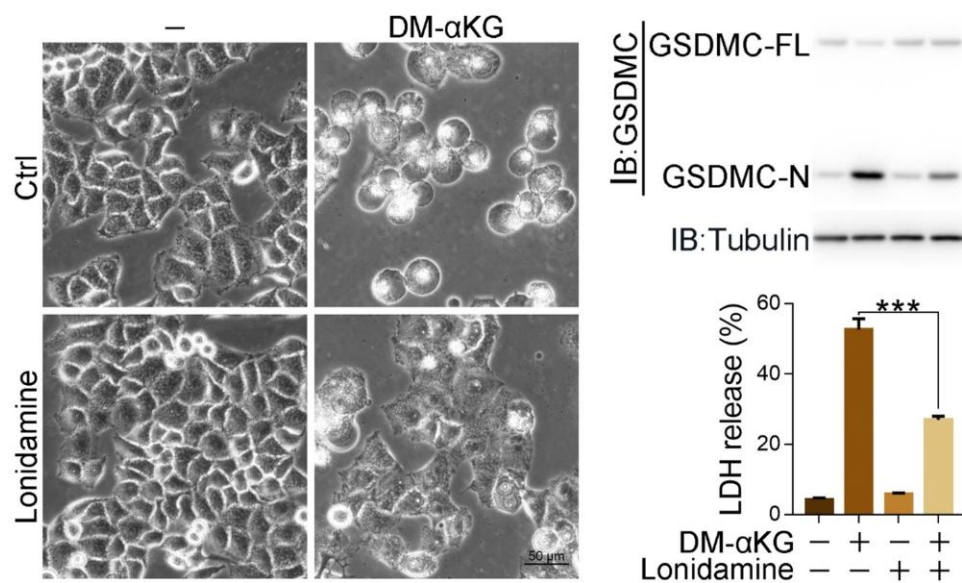
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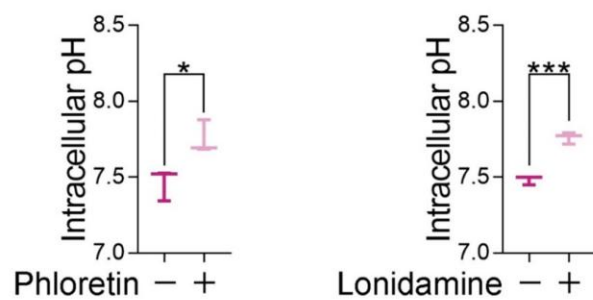
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h**i**

j



k

